## RECEIVED **CENTRAL FAX CENTER**

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## IN THE CLAIMS

212-318-3400

1. (currently amended) A method for the treatment of a skin disease comprising topically administering a subject in need thereof a therapeutically effective amount of a compound of formula (I) or a pharmaceutically acceptable salt thereof:

$$R^2$$
 $R^3$ 
 $R^4$ 
 $R^1$ 

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in which

R<sup>1</sup> is

(i) -C1-12-alkyl, straight-chain or branched-chain or -C2-C12 alkenyl, mono- or polyunsaturated,

optionally mono- or polysubstituted by -OH, -SH, -NH2, -NHC1-6-alkyl, -N( $C_{1-6}$ -alkyl)2, - $-NHC_{6-14}$  aryl,  $-N(C_{6-14}aryl)_2$ ,  $-N(C_{1-14}aryl)_2$   $-N(C_{1-6}alkyl)(C_{6-14}aryl)$ ,  $-NHCOR^6$ ,  $-NO_2$ , -CN, -F,  $-VC_{1-6}alkyl$ C1, -Br, -I, -O-C1-6-alkyl, -O-C6-14-aryl, -O(CO) $R^6$ , -S-C1-6-alkyl, -S-C6-14aryl, -SO $R^6$ , -SO<sub>3</sub>H, - $SO_2R^6, \ -OSO_2C_{1-6}-alkyl, \ -OSO_2C_{6-14}aryl, \ -(CS)R^6, \ -COOH, \ -(CO)R^6, \ mono-, \ bi \ or \ tricyclic$ saturated or mono- or polyunsaturated carbocycles having 3-14 ring members, mono-, bi- or

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tricyclic saturated or mono-or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, wherein the  $C_{6-14}$ aryl groups and the carbocyclic and heterocyclic substituents for their part can optionally be mono- or polysubstituted by  $R^4$ .

(ii) a mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycle having 3-14 ring members or a mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycle having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, or a carbo- or heterocyclic saturated or mono- or polyunsaturated spirocycle having 3-10 ring members, where heterocyclic systems contain 1-6 heteroatoms, which are preferably N, O and S, optionally mono- or polysubstituted by -OH, -SH, -NH<sub>2</sub>, NHC<sub>1-6</sub> alkyl, -N (C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, -N(C<sub>1-6</sub>alkyl) (C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup> -NO<sub>2</sub>, -CN, -F, -Cl, -Br, -I, -O-C-<sub>1-6</sub> alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>6</sup>, -OSO<sub>2</sub>C<sub>1-6</sub>alkyl, -OSO<sub>2</sub>C <sub>6-14</sub> aryl, -(CS)R<sup>6</sup>, -COOH, -(CO)R<sup>6</sup> mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members, mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, wherein the C<sub>6-14</sub>aryl groups and the carbocyclic and heterocyclic substituents can optionally be mono- or polysubstituted by R<sup>4</sup>,

R<sup>5</sup> is

a mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycle having 3-14 ring members or a mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycle having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, or a carbo-or heterocyclic saturated or mono-or polyunsaturated spirocycle having 3-10 ring members, where heterocyclic systems contain 1-6 heteroatoms, which preferably N, O and S, optionally mono- or polysubstituted by -OH, -SH, -NH<sub>2</sub> -NHC<sub>1-6</sub> alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>,

-N(C<sub>1-6</sub>alkyl)(C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup>, -NO<sub>2</sub> -CN, -F, -Cl, -Br, -I, -O-C-<sub>1.5</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1.6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>6</sup>, -OSO<sub>2</sub>C<sub>1-6</sub>alkyl, OSO<sub>2</sub>C<sub>6-14</sub>aryl, -(CS)R<sup>6</sup>, -COOH, -(CO)R<sup>6</sup>, mono-, bi or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members, mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S, wherein the C<sub>6-14</sub>aryl groups and the carbocyclic and heterocyclic substituents can optionally be mono- or polysubstituted by R<sup>4</sup> with the proviso that R<sup>5</sup> contains at least one substituent selected from -F, -Cl, -Br, -I;

 $R^2$ ,  $R^3$  are hydrogen or -OH, where at least one of the two substituents must be -OH;  $R^4$  is

-H, -OH, -SH, -NH<sub>2</sub> -NHC<sub>1-6</sub>-alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, -N(C<sub>1-6</sub> alkyl) (C<sub>6-14</sub>aryl), -NHCOR<sup>6</sup>, -NO<sub>2</sub>, -CN, -COOH, - (CO)R<sup>6</sup>, -(CS)R<sup>6</sup>, -F, --Cl, -Br, -I, -O-C<sub>1-6</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -O(CO)R<sup>6</sup>, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl, -SOR<sup>6</sup>, -SO<sub>2</sub>R<sup>6</sup>, -C<sub>1</sub>-C<sub>6</sub>-alkyl, wherein each aryl or alkyl may be mono- or polysubstituted by -OH, -F, -Cl, -Br, -I;

 $R^6$  is

-H, -NH<sub>2</sub>, -NHC<sub>1-6</sub>-alkyl, -N(C<sub>1-6</sub>-alkyl)<sub>2</sub>, -NHC<sub>6-14</sub>aryl, -N(C<sub>6-14</sub>aryl)<sub>2</sub>, - N(C<sub>1-6</sub>alkyl) (C<sub>6-14</sub>aryl), -O-C<sub>1-6</sub>-alkyl, -O-C<sub>6-14</sub>-aryl, -S-C<sub>1-6</sub>-alkyl, -S-C<sub>6-14</sub>aryl,

-C<sub>1-12</sub>-alkyl, straight-chain or branched-chain,

-C2-12-alkenyl, mono- or polyunsaturated, straight-chain or branched-chain,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members.

mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S;

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A is either a bond, or

-(CH<sub>2</sub>)<sub>m-</sub>, -(CH<sub>2</sub>)<sub>m</sub> -(CH=CH)<sub>n</sub> -(CH<sub>2</sub>)<sub>p</sub>-, -(CHOZ)<sub>m</sub>-, -(C=O)-, -(C=S)-, -(C=N-Z)-, -O-, -S-, -NZ-,

wherein m, p=0-3 and n=0-2 and

Z is

-H, or

-C1-12-alkyl, straight-chain or branched-chain,

-C2-12-alkenyl, mono- or polyunsaturated, straight-chain or branched-chain,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated carbocycles having 3-14 ring members,

mono-, bi- or tricyclic saturated or mono- or polyunsaturated heterocycles having 5-15 ring members and 1-6 heteroatoms, which are preferably N, O and S;

B is either carbon or sulfur, or -(S=O)-;

D is exygen sulfur O, S, CH2 or N-Z,

where, if B is carbon, D is O. S or CH2;

E is a bond, or

 $-(CH_2)_m$ -, -0-, -S-, -(N-Z)-, wherein m and Z have the meaning already described above.

2. (previously presented) The method of claim 1 wherein R<sup>5</sup> is selected from monocyclic saturated or mono- or polyunsaturated carbocycles and heterocycles having at least one halogen substituent.

- 3. (previously presented) The method of claim 2 wherein R<sup>5</sup> is selected from monocyclic aromatic carbocycles and heterocycles having at least one halogen substituent.
- 4. (previously presented) The method of claim 3 wherein R<sup>5</sup> is a pyridine ring having at least one halogen substituent.
- 5. (previously presented) The method of claim 3 wherein R<sup>5</sup> is a phenyl ring having at least one halogen substituent.
- 6. (previously presented) The method of claim 1 wherein  $R^1$  is selected from  $C_1$ - $C_{12}$  alkyl, which is optionally substituted.
- 7. (previously presented) The method of claim 1 wherein R<sup>1</sup> is selected from monocyclic saturated or mono- or polyunsaturated carbocycles or heterocycles, which are optionally substituted.
  - 8. (previously presented) The method of claim 1 wherein R<sup>2</sup> is OH and R<sup>3</sup> is H.
- 9. (previously presented) The method of claim 1 wherein A is selected from -(C=O)-and -(CHOH)-.
  - 10. (previously presented) The method of claim 1 wherein B is C.
  - 11. (previously presented) The method of claim 1 wherein D is O.
  - 12. (previously presented) The method of claim 1 wherein E is -(N--H)-.
- 13. (previously presented) The method of claim 1 wherein compound (I) is (N-3,5-dichloro-4-pyridinyl)-2-[1-(4-fluorobenzyl)-5-hydroxy-1 H-indol-3-yl]-2-oxoacetamide).
  - 14. (canceled)
  - 15. (previously presented) The method of claim 1 wherein the disease is an allergic.
  - 16. (canceled)

- 17. (previously presented) The method of claim 16 wherein the compound is administered to a skin area which is afflicted with the disease after an allergic challenge.
- 18. (previously presented) The method of claim 17 wherein the compound is administered up to 48 h after the allergic challenge.
  - 19. (canceled)
- 20. (previously presented) The method of claim 1 wherein a further pharmaceutical agent is administered and is a drug that stimulates cAMP production.
- 21. (previously presented) The method of claim 20 wherein the further pharmaceutical agent is a corticosteroid.
- 22. (previously presented) The method of claim 15, wherein the allergic disease is allergic dermatitis.
- 23. (previously presented) The method of claim 1, further comprising administering a further pharmaceutical agent.